



STIC Search Report

Biotech-Chem Library

STIC Database Tracking Number: 132662

TO: Shailendra Kumar
Location: 5c03 / 5c18
Tuesday, September 21, 2004
Art Unit: 1621
Phone: 272-0640
Serial Number: 10 / 706594

From: Jan Delaval
Location: Biotech-Chem Library
Rem 1A51
Phone: 272-2504

jan.delaval@uspto.gov

Search Notes

Jan please

Access DB#

132662

SEARCH REQUEST FORM

Scientific and Technical Information Center

Requester's Full Name: S. Kumar Examiner #: 69594 Date: 9/16/04
Art Unit: 1621 Phone Number 202-00640 Serial Number: 101706595
Mail Box and Bldg/Room Location: REM 5C18 Results Format Preferred (circle): PAPER DISK E-MAIL
5C03

If more than one search is submitted, please prioritize searches in order of need.

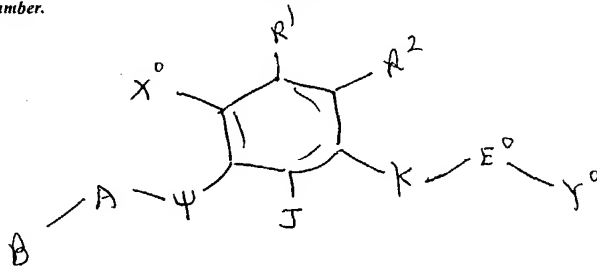
Please provide a detailed statement of the search topic, and describe as specifically as possible the subject matter to be searched. Include the elected species or structures, keywords, synonyms, acronyms, and registry numbers, and combine with the concept or utility of the invention. Define any terms that may have a special meaning. Give examples or relevant citations, authors, etc, if known. Please attach a copy of the cover sheet, pertinent claims, and abstract.

Title of Invention: Polycyclic aryl and heteroaryl substituted benzenes . . .

Inventors (please provide full names): Michael S. South et al.

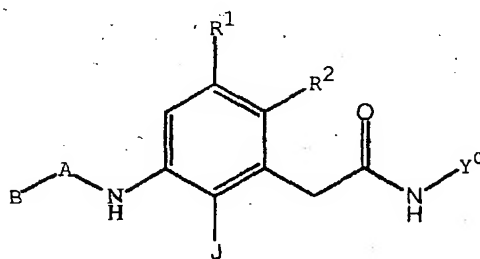
Earliest Priority Filing Date: 3/13/2000

For Sequence Searches Only Please include all pertinent information (parent, child, divisional, or issued patent numbers) along with the appropriate serial number.



See claim 1

Special



wherein;

[R² is 3-amino-5-carboxyphenyl, B is cyclobutyl, A is a bond, Y° is 4-amidinobenzyl, J is fluoro, and R¹ is hydrido;]

STAFF USE ONLY

Searcher: Jan
Searcher Phone #: 22504
Searcher Location: _____
Date Searcher Picked Up: 9/21
Date Completed: 9/21
Searcher Prep & Review Time: _____
Clerical Prep Time: 30
Online Time: 510

Type of Search

NA Sequence (#) _____
AA Sequence (#) _____
Structure (#) ☒ _____
Bibliographic _____
Litigation _____
Fulltext _____
Patent Family _____
Other _____

Vendors and cost where applicable

STN ☒ _____
Dialog _____
Questel/Orbit _____
Dr.Link _____
Lexis/Nexis _____
Sequence Systems _____
WWW/Internet _____
Other (specify) _____

=> fil reg

FILE 'REGISTRY' ENTERED AT 08:45:31 ON 21 SEP 2004
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STRUCTURE FILE UPDATES: 19 SEP 2004 HIGHEST RN 748118-51-6
DICTIONARY FILE UPDATES: 19 SEP 2004 HIGHEST RN 748118-51-6

TSCA INFORMATION NOW CURRENT THROUGH MAY 21, 2004

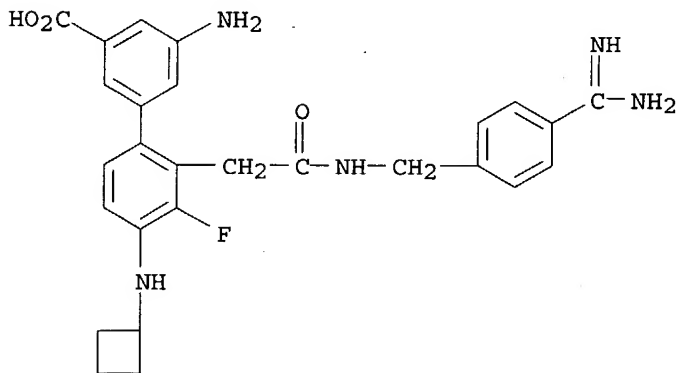
Please note that search-term pricing does apply when
conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. For more
information enter HELP PROP at an arrow prompt in the file or refer
to the file summary sheet on the web at:
<http://www.cas.org/ONLINE/DBSS/registryss.html>

=> d l3 ide can

L3 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2004 ACS on STN
RN 361340-15-0 REGISTRY
CN [1,1'-Biphenyl]-3-carboxylic acid, 5-amino-2'-[2-[[[4-
(aminoiminomethyl)phenyl]methyl]amino]-2-oxoethyl]-4'-(cyclobutylamino)-3'-
fluoro- (9CI) (CA INDEX NAME)
FS 3D CONCORD
MF C27 H28 F N5 O3
SR CA
LC STN Files: CA, CAPLUS, USPAT2, USPATFULL
DT.CA CAplus document type: Patent
RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES
(Uses)



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1907 TO DATE)
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 135:257039

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(FILE 'HOME' ENTERED AT 08:40:44 ON 21 SEP 2004)
SET COST OFF

FILE 'HCAPLUS' ENTERED AT 08:40:54 ON 21 SEP 2004
L1 1 S (US20040138275 OR US6660885 OR US20020025947)/PN OR (WO2001-U
SEL RN

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L2 381 S E1-E381
L3 1 S L2 AND C27H28FN5O3
SEL RN
L4 0 S E382/CRN

FILE 'HCAOLD' ENTERED AT 08:45:12 ON 21 SEP 2004
L5 0 S L3

FILE 'HCAPLUS' ENTERED AT 08:45:13 ON 21 SEP 2004
L6 1 S L3

FILE 'USPATFULL, USPAT2' ENTERED AT 08:45:13 ON 21 SEP 2004
L7 4 S L3

FILE 'REGISTRY' ENTERED AT 08:45:31 ON 21 SEP 2004

=> fil hcaplus

FILE 'HCAPLUS' ENTERED AT 08:46:36 ON 21 SEP 2004
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FILE COVERS 1907 - 21 Sep 2004 VOL 141 ISS 13
FILE LAST UPDATED: 20 Sep 2004 (20040920/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d l6 all hitstr

L6 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2004 ACS on STN
AN 2001:693283 HCAPLUS
DN 135:257039
ED Entered STN: 21 Sep 2001
TI Preparation of polycyclic aryl and heteroaryl substituted benzenes useful for selective inhibition of the coagulation cascade
IN South, Michael S.; Parlow, John J.
PA Pharmacia Corporation, USA
SO PCT Int. Appl., 437 pp.
CODEN: PIXXD2

DT Patent
 LA English
 IC ICM C07D211-26
 ICS C07D213-40; C07D521-00; C07C257-18; C07C311-21; A61K031-4465;
 A61P007-02
 CC 25-19 (Benzene, Its Derivatives, and Condensed Benzenoid Compounds)
 Section cross-reference(s): 1

FAN.CNT 1

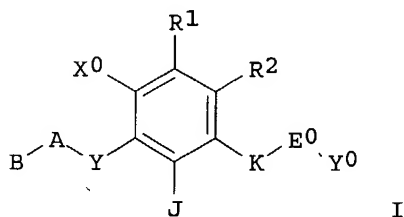
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|------|-----------------|--|----------|-----------------|----------|
| PI | WO 2001068605 | A1 | 20010920 | WO 2001-US7918 | 20010313 |
| | W: | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | |
| | RW: | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | |
| | AU 2001043598 | A5 | 20010924 | AU 2001-43598 | 20010313 |
| | US 2002025947 | A1 | 20020228 | US 2001-804959 | 20010313 |
| | US 6660885 | B2 | 20031209 | | |
| | US 2003236231 | A1 | 20031225 | US 2002-203866 | 20021113 |
| | US 2004138275 | A1 | 20040715 | US 2003-706595 | 20031112 |
| PRAI | US 2000-188943P | P | 20000313 | | |
| | US 2000-252159P | P | 20001120 | | |
| | US 2001-804959 | A1 | 20010313 | | |
| | WO 2001-US7918 | W | 20010313 | | |

CLASS

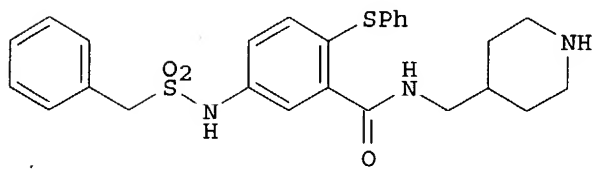
| PATENT NO. | CLASS | PATENT FAMILY CLASSIFICATION CODES |
|---------------|-------|---|
| WO 2001068605 | ICM | C07D211-26 |
| | ICS | C07D213-40; C07D521-00; C07C257-18; C07C311-21; A61K031-4465; A61P007-02 |
| US 2002025947 | ECLA | C07C257/18; C07C311/13; C07C323/49; C07C323/63; C07D211/26; C07D213/40B; C07D521/00B1N3 |

OS MARPAT 135:257039

GI



I



II

AB The title compds. [I; J = H, halo, OH, etc.; B = (un)substituted aryl, heteroaryl; A = a bond, CH₂SO₂, CH₂, (CH₂)₂, etc.; Y = NH, O, CO, etc.; X₀, R₁, R₂ = H, alkyl, halo, etc.; K = a bond, CH₂, etc.; E₀ = a bond, O, CONH, etc.; Y₀ = (4-piperidinyl)methyl, (amidino)benzyl, etc.] and their pharmaceutically acceptable salts, useful as inhibitors of serine proteases of the coagulation cascade, were prepared E.g., a multi-step synthesis of II.HCl which showed IC₅₀ of > 30 μ M against factor VIIa, factor Xa and thrombin, and IC₅₀ of 0.3 μ M against trypsin, was given.

ST benzamidine polycyclic aryl heteroaryl prepn anticoagulant platelet aggregation inhibitor; benzene polycyclic aryl heteroaryl prepn anticoagulant platelet aggregation inhibitor; serine protease inhibitor benzamidine polycyclic aryl heteroaryl prepn; factor VIIa Xa inhibitor benzamidine polycyclic aryl heteroaryl prepn; thrombin inhibitor benzamidine polycyclic aryl heteroaryl prepn; trypsin inhibitor benzamidine polycyclic aryl heteroaryl prepn

IT Anticoagulants
Platelet aggregation inhibitors
(preparation of polycyclic aryl and heteroaryl substituted benzenes useful for selective inhibition of the coagulation cascade)

IT

| | | | | |
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 361339-75-5P 361339-76-6P 361339-77-7P 361339-78-8P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of polycyclic aryl and heteroaryl substituted benzenes useful for selective inhibition of the coagulation cascade)

IT 361339-79-9P 361339-80-2P 361339-81-3P 361339-82-4P 361339-83-5P
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 361340-48-9P 361340-49-0P 361340-50-3P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of polycyclic aryl and heteroaryl substituted benzenes useful for selective inhibition of the coagulation cascade)

IT 9002-04-4, thrombin 9002-05-5, factor Xa 9002-07-7, trypsin
 37259-58-8, serine protease 65312-43-8, factor VIIa

RL: BSU (Biological study, unclassified); MSC (Miscellaneous); BIOL (Biological study)

(preparation of polycyclic aryl and heteroaryl substituted benzenes useful for selective inhibition of the coagulation cascade)

IT 67-64-1, Acetone, reactions 100-52-7, Benzaldehyde, reactions
 108-98-5, Thiophenol, reactions 122-78-1, Phenylacetaldehyde 591-19-5,
 m-Bromoaniline 813-19-4, Bis(tributyltin) 960-16-7 1939-99-7,
 α -Toluenesulfonyl chloride 3731-53-1, 4-Pyridylmethylamine
 5036-48-6, 3-(Imidazol-1-yl)propylamine 7144-05-0, 4-Piperidinemethanamine
 7304-32-7, 2-Fluoro-5-nitrobenzoic acid 19293-58-4, 4-Dimethylaminobenzylamine
 85068-28-6, 2,6-Difluorophenylacetic acid 147291-50-7 162698-22-8 226942-83-2
 361337-21-5 361337-22-6

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of polycyclic aryl and heteroaryl substituted benzenes useful for selective inhibition of the coagulation cascade)

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 361337-19-1P 361337-20-4P 361340-51-4P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of polycyclic aryl and heteroaryl substituted benzenes useful for selective inhibition of the coagulation cascade)

IT 361336-87-0P 361336-91-6P 361337-07-7P

RL: SPN (Synthetic preparation); PREP (Preparation)

(preparation of polycyclic aryl and heteroaryl substituted benzenes useful for selective inhibition of the coagulation cascade)

RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE

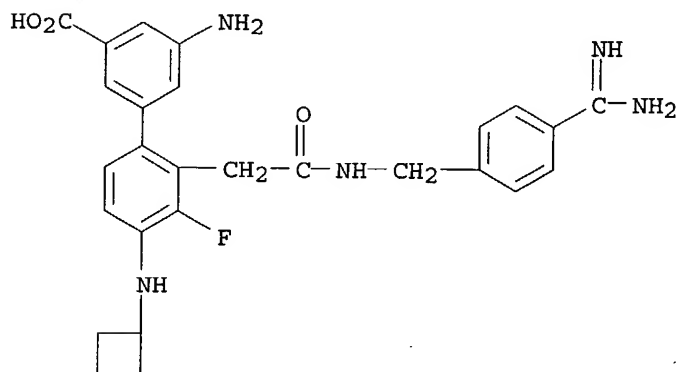
- (1) Illig, C; US 5741819 A 1998 HCAPLUS
- (2) Ljungberg; EUR J PHAR SCI 2001, V12(4), P441 HCAPLUS
- (3) Terumo Corp; JP 07233148 A 1995 HCAPLUS

IT 361340-15-0P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of polycyclic aryl and heteroaryl substituted benzenes useful for selective inhibition of the coagulation cascade)

RN 361340-15-0 HCAPLUS

CN [1,1'-Biphenyl]-3-carboxylic acid, 5-amino-2'-[2-[[[4-(aminoiminomethyl)phenyl]methyl]amino]-2-oxoethyl]-4'-(cyclobutylamino)-3'-fluoro- (9CI) (CA INDEX NAME)



=> fil uspatall

FILE 'USPATFULL' ENTERED AT 08:46:44 ON 21 SEP 2004

CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'USPAT2' ENTERED AT 08:46:44 ON 21 SEP 2004

CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

=> d bib abs hitstr tot.l7

L7 ANSWER 1 OF 4 USPATFULL on STN

AN 2004:179105 USPATFULL

TI Polycyclic aryl and heteroaryl substituted benzenes useful for selective inhibition of the coagulation cascade

IN South, Michael S., St. Louis, MO, UNITED STATES

Parlow, John J., Arnold, MO, UNITED STATES

PA Pharmacia Corporation (U.S. corporation)

PI US 2004138275 A1 20040715

AI US 2003-706595 A1 20031112 (10)

RLI Continuation of Ser. No. US 2001-804959, filed on 13 Mar 2001, GRANTED, Pat. No. US 6660885

PRAI US 2000-252159P 20001120 (60)

US 2000-188943P 20000313 (60)

DT Utility

FS APPLICATION

LREP SENNIGER POWERS LEAVITT AND ROEDEL, ONE METROPOLITAN SQUARE, 16TH FLOOR, ST LOUIS, MO, 63102

CLMN Number of Claims: 59

ECL Exemplary Claim: 1

DRWN No Drawings

LN.CNT 12687

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The invention relates to polycyclic aryl and heteroaryl substituted benzene compounds useful as inhibitors of serine proteases of the coagulation cascade and compounds, compositions and methods for anticoagulant therapy for the treatment and prevention of a variety of thrombotic conditions including coronary artery and cerebrovascular diseases.

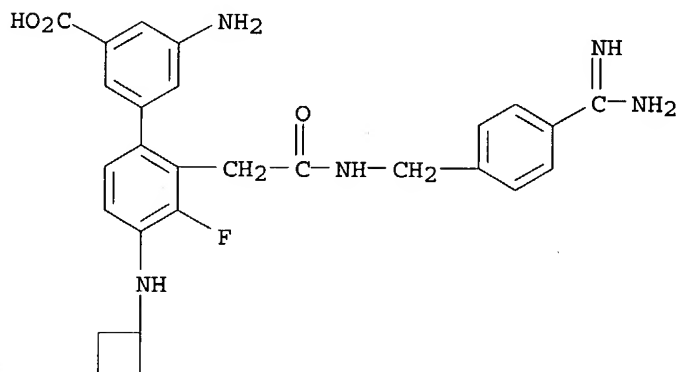
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 361340-15-0P

(preparation of polycyclic aryl and heteroaryl substituted benzenes useful for selective inhibition of the coagulation cascade)

RN 361340-15-0 USPATFULL

CN [1,1'-Biphenyl]-3-carboxylic acid, 5-amino-2'-[2-[[[4-(aminoiminomethyl)phenyl]methyl]amino]-2-oxoethyl]-4'-(cyclobutylamino)-3'-fluoro- (9CI) (CA INDEX NAME)



L7 ANSWER 2 OF 4 USPATFULL on STN

AN 2003:335353 USPATFULL

TI Polycyclic aryl and heteroaryl substituted benzenes useful for selective inhibition of the coagulation cascade

IN South, Michael S, St Louis, MO, UNITED STATES

Parlow, John J, Arnold, MO, UNITED STATES

PI US 2003236231 A1 20031225

AI US 2002-203866 A1 20021113 (10)

WO 2001-US7918 20010313

DT Utility

FS APPLICATION

LREP SENNIGER POWERS LEAVITT AND ROEDEL, ONE METROPOLITAN SQUARE, 16TH FLOOR, ST LOUIS, MO, 63102

CLMN Number of Claims: 59

ECL Exemplary Claim: 1

DRWN No Drawings

LN.CNT 12689

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The invention relates to polycyclic aryl and heteroaryl substituted benzene compounds useful as inhibitors of serine proteases of the coagulation cascade and compounds, compositions and methods for anticoagulant therapy for the treatment and prevention of a variety of thrombotic conditions including coronary artery and cerebrovascular diseases.

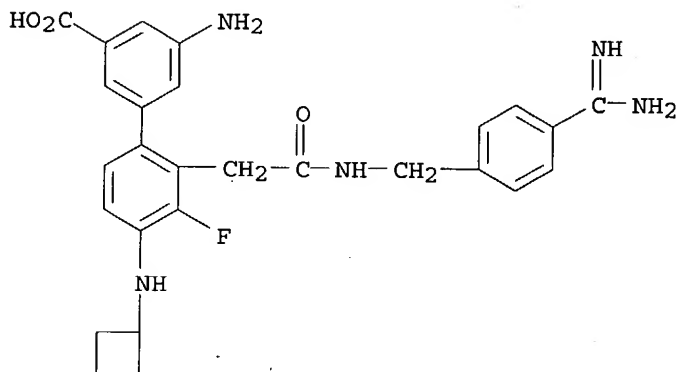
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 361340-15-0P

(preparation of polycyclic aryl and heteroaryl substituted benzenes useful for selective inhibition of the coagulation cascade)

RN 361340-15-0 USPATFULL

CN [1,1'-Biphenyl]-3-carboxylic acid, 5-amino-2'-[2-[[[4-(aminoiminomethyl)phenyl]methyl]amino]-2-oxoethyl]-4'-(cyclobutylamino)-3'-fluoro- (9CI) (CA INDEX NAME)



L7 ANSWER 3 OF 4 USPATFULL on STN

AN 2002:43580 USPATFULL

TI Polycyclic aryl and heteroaryl substituted benzenes useful for selective inhibition of the coagulation cascade

IN South, Michael S., St. Louis, MO, UNITED STATES
Parlow, John J., Arnold, MO, UNITED STATES

PI US 2002025947 A1 20020228

US 6660885 B2 20031209

AI US 2001-804959 A1 20010313 (9)

PRAI US 2000-252159P 20001120 (60)

US 2000-188943P 20000313 (60)

DT Utility

FS APPLICATION

LREP SENNIGER POWERS LEAVITT AND ROEDEL, ONE METROPOLITAN SQUARE, 16TH FLOOR,
ST LOUIS, MO, 63102

CLMN Number of Claims: 59

ECL Exemplary Claim: 1

DRWN No Drawings

LN.CNT 12668

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The invention relates to polycyclic aryl and heteroaryl substituted benzene compounds useful as inhibitors of serine proteases of the coagulation cascade and compounds, compositions and methods for anticoagulant therapy for the treatment and prevention of a variety of thrombotic conditions including coronary artery and cerebrovascular diseases.

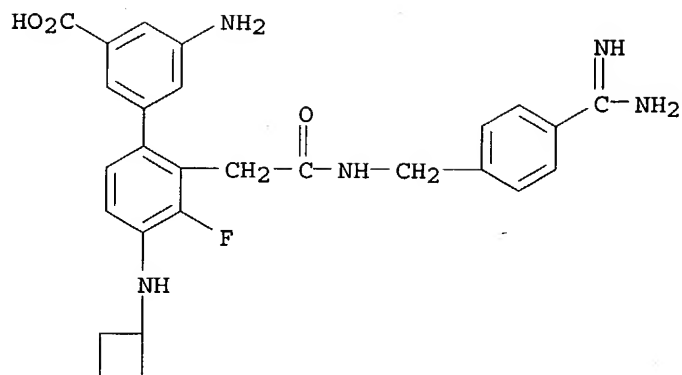
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 361340-15-0P

(preparation of polycyclic aryl and heteroaryl substituted benzenes useful for selective inhibition of the coagulation cascade)

RN 361340-15-0 USPATFULL

CN [1,1'-Biphenyl]-3-carboxylic acid, 5-amino-2'-[2-[[[4-(aminoiminomethyl)phenyl]methyl]amino]-2-oxoethyl]-4'-(cyclobutylamino)-3'-fluoro- (9CI) (CA INDEX NAME)



L7 ANSWER 4 OF 4 USPAT2 on STN
 AN 2002:43580 USPAT2
 TI Polycyclic aryl and heteroaryl substituted benzenes useful for selective inhibition of the coagulation cascade
 IN South, Michael S., St. Louis, MO, United States
 Parlow, John J., Arnold, MO, United States
 PA Pharmacia Corporation, Skokie, IL, United States (U.S. corporation)
 PI US 6660885 B2 20031209
 AI US 2001-804959 20010313 (9)
 PRAI US 2000-252159P 20001120 (60)
 US 2000-188943P 20000313 (60)
 DT Utility
 FS GRANTED
 EXNAM Primary Examiner: Kumar, Shailendra
 CLMN Number of Claims: 26
 ECL Exemplary Claim: 1
 DRWN 0 Drawing Figure(s); 0 Drawing Page(s)
 LN.CNT 8147
 CAS INDEXING IS AVAILABLE FOR THIS PATENT.
 AB The invention relates to polycyclic aryl and heteroaryl substituted benzene compounds useful as inhibitors of serine proteases of the coagulation cascade and compounds, compositions and methods for anticoagulant therapy for the treatment and prevention of a variety of thrombotic conditions including coronary artery and cerebrovascular diseases.

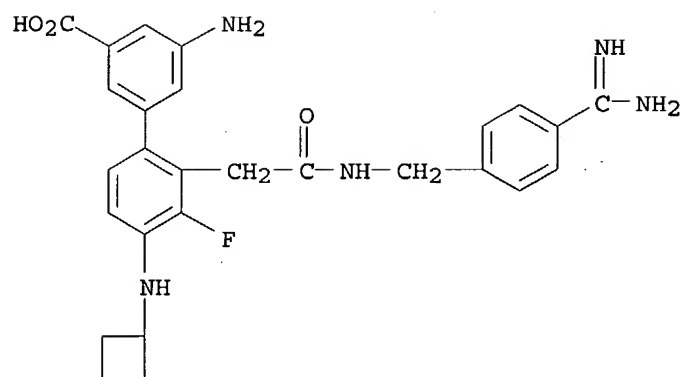
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 361340-15-0P

(preparation of polycyclic aryl and heteroaryl substituted benzenes useful for selective inhibition of the coagulation cascade)

RN 361340-15-0 USPAT2

CN [1,1'-Biphenyl]-3-carboxylic acid, 5-amino-2'-[2-[[[4-(aminoiminomethyl)phenyl]methyl]amino]-2-oxoethyl]-4'-(cyclobutylamino)-3'-fluoro- (9CI) (CA INDEX NAME)



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